June 19, 2003

UNITED STATES PATENT AND TRADEMARK OFFICE

Examiner: To Be Assigned (Rita Desai in parent appl. No. 10/126,506)

Art Unit: To Be Assigned (1614 in parent appl. No. 10/126,506)

Re: Application of: R. Richard GOEHRING, et al.

Serial No.: To Be Assigned

Filed: Herewith

For: SPIROPYRAZOLE COMPOUNDS

INFORMATION DISCLOSURE STATEMENT

MAIL STOP:

Commissioner for Patents P.O. Box 1450

Alexandria, VA 22313-1450

Sir:

In accordance with the provisions of 37 C.F.R. § 1.97, Applicants hereby make of record the references listed on the accompanying Form PTO-1449 (3 pages) for consideration by the Examiner in connection with the examination of the above-identified patent application. These references were provided to the U. S. Patent and Trademark Office in the Information Disclosure Statement filed March 11, 2003 for the parent application, Patent Application No. 10/126,506.

Applicants respectfully direct the Examiner's attention to reference BK listed on the accompanying Form PTO-1449 which was cited in the International Search Report issued by the PCT for the corresponding International Application PCT/US02/12376. It is noted that references JP 60-120872 (reference BI) and JP 10-330377 (reference BJ) are directed to the chemical structures shown therein and are readily understandable.

This Information Disclosure Statement is being filed under 37 C.F.R. §1.97 (b), before the mailing date of a First Office Action on the merits. Accordingly, no fee is believed to be due

for submitting this Information Disclosure Statement. In the event any fee is due, the Assistant Commissioner is authorized to charge the deficiency to our Deposit Account No. 50-0552.

Applicants respectfully request that the references cited in the accompanying Form PTO-1449 be considered and made of record. If any of the listed references are found to be missing, the Examiner is requested to contact the undersigned so that a copy may be promptly forwarded.

Respectfully submitted,

DAVIDSON, DAVIDSON & KAPPEL, LLC

 $\mathbf{R}_{\mathbf{V}}$

Robert J. Paragiso

Reg. No. 41,24

Davidson, Davidson & Kappel, LLC 485 Seventh Avenue, 14th Floor New York, New York 10018 (212) 736-2427

FORM PTO-1449 U.S. DEPARTMENT OF COMMERCE ATTY, DOCKET NO .: SERIAL NO.: 10/126,506 (REV. 7-80) PATENT AND TRADEMARK OFFICE 332.1132 LIST OF PRIOR ART CITED BY APPLICANT APPLICANT(S): R. Richard GOEHRING, et al. (Use several sheets if necessary) FILING DATE: April 18, 2002 GROUP: 1614 U.S. PATENT DOCUMENTS *EXAMINER DOCUMENT NUMBER CLASS DATE NAME **SUBCLASS** FILING DATE INITIAL IF APPROPRIATE AA 12/15/64 Paul Adriaan, et al. 260 293.4 3 3 9 AΒ 8 0 0 05/09/67 Paul Adriaan, et al. 260 294 AC 3 3 2 5 4 9 9 06/13/67 260 294 Poos, et al. ΑD 2 5 07/29/80 424 248.5 Mentrup, et al. ΑE 4 3 2 9 3 3 05/11/82 Stokbroekx, et al. 424 267 4 ΑF 4 0 5 2 8 10/18/83 Teranishi, et al. 424 251 5 7 AG 5 4 0 4 4 11/12/96 316 Thompson, et al. 514 5 6 9 6 6 08/26/97 Di Malta, et al. AH 514 387 7 ΑI 0 0 5 4 06/02/98 Huff, et al. 6 514 302 ΑJ 5 7 6. 7 8 06/16/98 Nargund, et al. 514 226.4 7 6 0 9 ΑK 6 3 6 05/16/00 Yang, et al. 514 322 6 0 3 7 ΑL 12/26/00 Budhu, et al. 514 326 FOREIGN PATENT DOCUMENTS DOCUMENT NUMBER COUNTRY CLASS SUBCLASS TRANSLATION DATE YES NO 9 AM 0 3 03/20/97 WO (A1) C07D 211/44 Abstract 9 7 0 3 10/30/97 AN 4 n 5 WO (A1) C07D 401/04 9 8 12/03/98 ΑO 8 WO (A1) C07D 401/04 6 Abstract AP 9 9 2 9 6 9 6 06/17/99 WO (A1) C07D 491/107 9 9 AQ 3 6 4 2 07/22/99 WO (A1) C07D 401/04 AR 9 9 9 4 2 09/30/99 WO (A1) 31/165 A61K Abstract OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.) AS Custers, F.G.J., et al., "Vesamicol and some of its derivatives: Questionable ligands for selectively labelling acetylcholine transporters in rat brain", European Journal of Pharmacology, 338: 117-183, 1997. **EXAMINER** DATE CONSIDERED *EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

FORM PTO-1449 U.S. DEPARTMENT OF COMMERCE ATTY. DOCKET NO .: SERIAL NO.: 10/126,506 (REV. 7-80) PATENT AND TRADEMARK OFFICE 332.1132 APPLICANT(S): R. Richard GOEHRING, et al. LIST OF PRIOR ART CITED BY APPLICANT (Use several sheets if necessary) GROUP: 1614 FILING DATE: April 18, 2002 U.S. PATENT DOCUMENTS **CLASS SUBCLASS** *EXAMINER DOCUMENT NUMBER DATE NAME FILING DATE INITIAL IF APPROPRIATE 01/09/01 252.13 BA Ito, et al. 514 FOREIGN PATENT DOCUMENTS DOCUMENT NUMBER DATE COUNTRY CLASS **SUBCLASS** TRANSLATION YES NO BB 9 9 9 9 9 7 11/25/99 WO (A1) C07D 471/10 5 5 0 0 0 4 02/10/00 WO (A1) C07D 211/52 BC 6 BD n 0 7 0 5 0 02/01/01 WO (A1) A61K 31/445 4 2 06/21/01 C07D 265/18 BE 0 1 4 1 3 WO (A1) 2 BF 0 2 0 0 03/14/02 WO (A2) A61K 31/347 0 9 9 10/26/83 C07D 401/06 BG 0 EP (A2) вн 0 9 2 1 1 2 5 06/09/99 EP (A1) C07D 471/10 2 0 2 06/28/85 C07D 237/34 ВΙ 60 1 8 Japan 7 BJ 10 3 0 12/15/98 C07D 401/14 Japan OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.) Harrison T, et al., "High Affinity, Selective Neurokinin 2 and Neurokinin 3 Receptor Antagonists from a Common Structural BK Template," Bioorganic & Medicinal Chemistry Letters., Vol. 8, pp 1343-1348, especially compounds 2d and 3d on page 1345. (1998). BL. Takai H, et al., Synthesis and Pharmacological Evaluation of Peperidine Derivatives with Various Heterocyclic Rings at the 4-Position, Chemical Pharmaceutical Bulletin, March 1985, Vol. 33, No. 3, pages 1105-1115, especially pages 1105 and 1106 BM Database on STN CASDATA (Columbus, Ohio, USA) CA accession: no. 129:54361, Huff, et al., 'Preparation of benzisothiazolones and analogs as alpha. 1C. adrenergic receptor antagonist', 1998, 392146 CAPLUS (Abstract). BN Database on STN CASDATA (Columbus, Ohio, USA) CA accession: no. 130:95479, Sasaki, et al., 'Preparation of piperidine derivatives as cell adhesion inhibitors for inflammation inhibitors, metastasis inhibitors, etc.', 1998, 795478 CAPLUS (Abstract). Database on STN CADATA (Columbus, Ohio, USA) CA accession: no. 124:176079, Huff, et al., 'Preparation of heterocycles as alpha-BO 1c adrenergic receptor antagonists', 1995, 998362 CAPLUS (Abstract). BP Database on STN CASDATA (Columbus, Ohio, USA) CA accession: no. 81:77786, Klein, et al., 'Potential analgesics. 3. 1-(4-Piperidinyl) -2-indolinones and -3, 4-dihydrocarbostyrils', Arch. Pharm. Vol. 307 (5), pp 360-365. (1974), 1974, 47786 CAPLUS (Abstract). BQ Database on STN CASDATA (Columbus, Ohio, USA) CA accession: no. 136:53656, Forbes I.T., 'A short and efficient synthesis of N-substituted indol-2-ones (oxindoles)', Tetrahedron Let., (2001), vol 42 (39), pp. 6943 - 6945, 2001:674545 CAPLUS (Abstract). **EXAMINER** DATE CONSIDERED *EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

FORM PTO-14- (REV. 7-80)	U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE								ATTY. DOCKET NO.: 332.1132		SERIAL NO.: 10/126,506			
LIST OF PRIOR ART CITED BY APPLICANT									APPLICANT(S): R. Richard GOEHRING, et al.					
(Use several sheets if necessary)								FILING DATE:	April 18, 2002	GROUP: 1614				
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	Cl	Data diph	abase o	n STN kylpip	N CASI	DATA deriva	(Colur	nbus, (Ohio, USA) CA	A accession: no. 13: receptor agoinsts',	5:272955, Tsushi 2001:713309 CA	ma, et al., 'Prepar PLUS (Abstract).	ation of	
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PATENT COOPERATION TREATY

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INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference 332.1132PCT	FOR FURTHER see Notification of Transmittal of International Search ACTION (Form PCT/ISA/220) as well as, where applicable, item								
International application No.	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)							
PCT/US02/12376	18 APRIL 2002	18 APRIL 2001							
Applicant EURO-CELTIQUE, S.A.									
	n prepared by this International Searching Aug transmitted to the International Bureau.	thority and is transmitted to the applicant							
X It is also accompanied by a c	opy of each prior art document cited in this	report.							
1. Basis of the report		•							
, , , , , , , , , , , , , , , , , , , ,	he international search was carried out on the	pasis of the international application in the							
	unless otherwise indicated under this item. carried out on the basis of a translation of the	ne international application furnished to this							
b. With regard to any nucleotide was carried out on the basis of	and/or amino acid sequence disclosed in the in the sequence listing:	nternational application, the international search							
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2. Certain claims were found	unsearchable (See Box I).								
5. Unity of invention is lacking	ng (See Box II).								
4. With regard to the title,									
X the text is approved as sub	mitted by the applicant.								
the text has been established	ed by this Authority to read as follows:								
5. With regard to the abstract,	· <u>.</u>								
X the text is approved as sub	mitted by the applicant.								
Box III. The applicant may,	the text has been established, according to Rule 38.2(b), by this Authority as it appears in Box III. The applicant may, within one month from the date of mailing of this international search report, submit comments to this Authority.								
6. The figure of the drawings to be p	ublished with the abstract is Figure No.								
as suggested by the applica	as suggested by the applicant.								
because the applicant failed	to suggest a figure.	None of the figures.							
because this figure better c									

INTERNATIONAL SEARCH REPORT

International application No. PCT/US02/12376

A. CLASSIFICATION OF SUBJECT MATTER								
IPC(7) :A61K 31/438; C07D 471/10								
US CL :514/278; 546/20 According to International Patent Classification (IPC) or to both national classification and IPC								
B. FIELDS SEARCHED								
Minimum documentation searched (classification system followed by classification symbols)								
U.S.: 514/278; 546/20								
0.0. : 514/2/6;546/20								
Documentation searched other than minimum documentation to the extent that such documents are included in the	ields							
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Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)								
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C. DOCUMENTS CONSIDERED TO BE RELEVANT								
Category* Citation of document, with indication, where appropriate, of the relevant passages Relevant to	claim No.							
A HARRISON, T. et al. High Affinity, Selective Neurokinin 2 and 1-31								
Neurokinin 3 Receptor Antagonists from a common structural								
template. Bioorg. & Medicin. Chem. Lett. 1998, Vol. 8, pages								
1343-1348, especially compounds 2d and 3d on page 1345.								
	5							
Further documents are listed in the continuation of Box C. See patent family annex.								
"T" later document published after the international filing date and not in conflict with the application but cited to								
"A" document defining the general state of the art which is not considered the principle or theory underlying the invention to be of particular relevance								
"E" earlier document published on or after the international filing date considered novel or cannot be considered to involve an in-								
"L" document which may throw doubts on priority claim(s) or which is when the document is taken alone cited to establish the publication date of another citation or other	•							
special reason (as specified) "Y" document of particular relevance; the claimed invention considered to involve an inventive step when the document	document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined							
"O" document referring to an oral disclosure, use, exhibition or other with one or more other such documents, such combined means obvious to a person skilled in the art								
"P" document published prior to the international filing date but later "&" document member of the same patent family than the priority date claimed								
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Name and mailing address of the ISA/US Authorized officer 12 and 100	Authority officer Dudgles for							